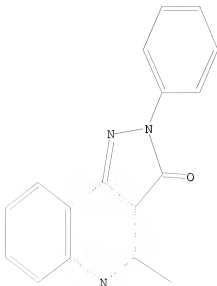


L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



G1 H,Ak

Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 15:50:01 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 88 TO ITERATE

100.0% PROCESSED 88 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 1198 TO 2322

PROJECTED ANSWERS: 2 TO 124

L2 2 SEA SSS SAM L1

=> s l1 ful

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100.0% PROCESSED 1687 ITERATIONS

30 ANSWERS

SEARCH TIME: 00.00.01

L3 30 SEA SSS FUL L1

10537538

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COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	178.36	178.57

FILE 'CAPLUS' ENTERED AT 15:50:14 ON 22 SEP 2008
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FILE COVERS 1907 - 22 Sep 2008 VOL 149 ISS 13
FILE LAST UPDATED: 21 Sep 2008 (20080921/ED)

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L4 2 L3

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L4 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2008 ACS on STN
GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The invention relates to a preparation of novel pyrazolo[4,3-c]quinoline derive. of formula I [wherein: Z is carboxylic acid or ester group; X is a bond or a divalent group selected from alkylene, or NHC(O), etc.; Y is O, S, N-oxide, or NH, etc.; R1 and R3 are independently selected from H, F, Cl, NO2, or CN, etc.; R2 (un)substituted C3-7cycloalkyl or phenyl] as immunomodulatory agents, useful for the treatment of rheumatoid arthritis, multiple sclerosis, diabetes, asthma, transplantation, systemic lupus erythematosis, and psoriasis. The title compds. are CD80 antagonists capable of inhibiting the interactions between CD80 and CD28. For instance, IC50 for the prepared pyrazoloquinoline derivative II was 3.4 μ M

(example 7).

AN 2004:534202 CAPLUS

DN 141:71540

TI A preparation of pyrazolo[4,3-c]quinoline derivatives, useful as immunomodulatory agents

IN Matthews, Ian Richard; Huxley, Philip; Magaraci, Filippo; Brennan, Chris James; Uddin, Muhammed Kamal; Pettersson, Lars Olof Goeran; Thrige, Dortha Da Graca

PA Active Biotech Ab, Swed.

SO PCT Int. Appl., 28 pp.

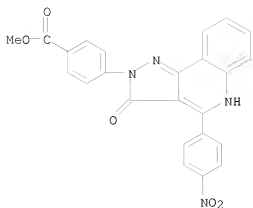
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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PI	WO 2004055014	A1	20040701	WO 2003-SE1941	20031212
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	EP 1572689	A1	20050914	GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK	
	US 20060035919	A1	20060216	SE 2002-3722 US 2002-433580P WO 2003-SE1941 US 2005-537538 SE 2002-3722 US 2002-433580P WO 2003-SE1941	A 20021216 P 20021216 W 20031212 20050915 A 20021216 P 20021216 W 20031212
OS	MARPAT 141:71540				
IT	713141-32-3P				
	RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)				
	(preparation of pyrazoloquinoline derivs. useful as immunomodulatory agents)				
RN	713141-32-3 CAPLUS				
CN	Benzoic acid, 4-[3,5-dihydro-4-(4-nitrophenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]-, methyl ester (CA INDEX NAME)				

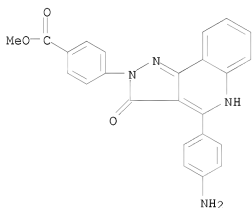


IT 713141-33-4P 713141-34-5P 713141-35-6P
 713141-36-7P 713141-37-8P 713141-38-9P
 713141-39-0P 713141-40-3P 713141-41-4P
 713141-42-5P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
 (Uses)
 (preparation of pyrazoloquinoline derivs. useful as immunomodulatory agents)

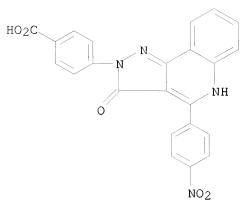
RN 713141-33-4 CAPLUS

CN Benzoic acid, 4-[4-(4-aminophenyl)-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-
 c]quinolin-2-yl]-, methyl ester (CA INDEX NAME)



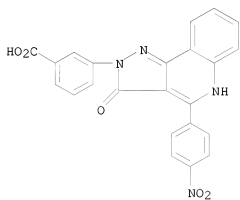
RN 713141-34-5 CAPLUS

CN Benzoic acid, 4-[3,5-dihydro-4-(4-nitrophenyl)-3-oxo-2H-pyrazolo[4,3-
 c]quinolin-2-yl]- (CA INDEX NAME)



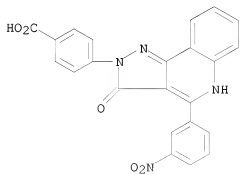
RN 713141-35-6 CAPLUS

CN Benzoic acid, 3-[3,5-dihydro-4-(4-nitrophenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]- (CA INDEX NAME)



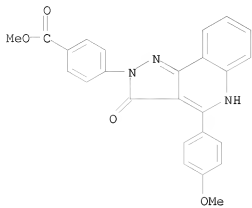
RN 713141-36-7 CAPLUS

CN Benzoic acid, 4-[3,5-dihydro-4-(3-nitrophenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]- (CA INDEX NAME)



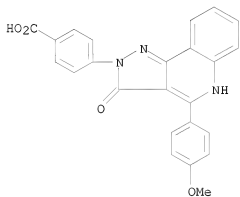
RN 713141-37-8 CAPLUS

CN Benzoic acid, 4-[3,5-dihydro-4-(4-methoxyphenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]-, methyl ester (CA INDEX NAME)



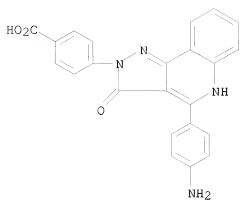
RN 713141-38-9 CAPLUS

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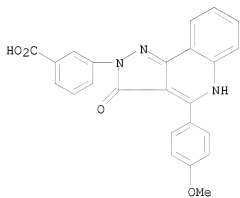
RN 713141-39-0 CAPLUS

CN Benzoic acid, 4-[4-(4-aminophenyl)-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]- (CA INDEX NAME)



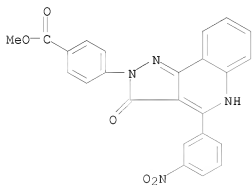
RN 713141-40-3 CAPLUS

CN Benzoic acid, 3-[3,5-dihydro-4-(4-methoxyphenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]- (CA INDEX NAME)



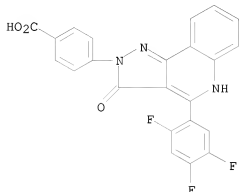
RN 713141-41-4 CAPLUS

CN Benzoic acid, 4-[3,5-dihydro-4-(3-nitrophenyl)-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl]-, methyl ester (CA INDEX NAME)



RN 713141-42-5 CAPLUS

CN Benzoic acid, 4-[3,5-dihydro-3-oxo-4-(2,4,5-trifluorophenyl)-2H-pyrazolo[4,3-c]quinolin-2-yl]- (CA INDEX NAME)

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMATL4 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2008 ACS on STN
GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The invention relates to novel heterocyclic compds., to methods for their preparation, to compns. containing them, and to methods and use for clin. treatment

of medical conditions which may benefit from immunomodulation, including rheumatoid arthritis, multiple sclerosis, diabetes, asthma, transplantation, systemic lupus erythematosus, and psoriasis. More particularly, the invention relates to novel heterocyclic compds. I, which are CD80 antagonists capable of inhibiting the interactions between CD80 and CD28. In formula I, R1 and R3 independently represent H, F, Cl, Br, NO₂, CN, C1-C6 alkyl optionally substituted by F or Cl, or C1-C6 alkoxy optionally substituted by F; R2 represents H, or optionally substituted C1-C6 alkyl, C3-C7 cycloalkyl, or optionally substituted Ph; Y represents O, S, N-oxide, or N(R5), wherein R5 represents H or C1-C6 alkyl; X represents a bond or a divalent C1-C6 alkylene radical; R4 represents -C(O)NR6R7, -NR7C(O)OR6, -NR7C(O)OR6, -NHC(O)NHR6, or -NHC(S)NHR6, wherein R6 represents H, or a radical of formula -(Alk)b-Q wherein b = 0-1 and Alk is an optionally substituted divalent straight chain or branched C1-C12 alkylene, C2-C12 alkenylene or C2-C12 alkynylene radical which may be interrupted by one or more non-adjacent -O-, -S- or -N(R8)- radicals wherein R8 represents H or C1-C4 alkyl, C3-C4 alkenyl, C3-C4 alkynyl, or C3-C6 cycloalkyl, and Q represents H, CF₃, OH, SH, NR8R8 wherein each R8 may be the same or different, an ester group, or an optionally substituted Ph, C3-C7 cycloalkyl, C5-C7 cycloalkenyl or heterocyclic ring having from 5 to 8 ring atoms; and R7 represents H or C1-C6 alkyl; or when taken

together with the atom or atoms to which they are attached, R6 and R7 form an optionally substituted heterocyclic ring having from 5 to 8 ring atoms. Approx. 170 example compds. and several intermediates were prepared. For instance, invention compound II (claimed individually) was prepared in 5 steps: (1) cyclocondensation of 3-cyclopropyl-3-oxopropionic acid Me ester with Et 2-aminobenzoate to give a quinolone derivative, (2) conversion of the quinolone ester to a chloroquinoline ester with PCCl₃, (3) cyclocondensation of the latter with 4-hydrazinobenzoic acid to form the pyrazole ring, (4) conversion of the free acid group to an acid chloride, and (5) amidation with H₂N(CH₂)₃NMe₂. In a cell-free, Eu/APC-based, homogeneous time-resolved fluorescence (HTRF) assay, used to determine inhibition of CD80-CD28 interaction, II had EC₅₀ < 1 μM.

AN 2004:467892 CAPLUS

DN 141:38606

TI Pyrazoloquinolines and analogs with CD80 antagonist immunomodulating activity, and their preparation, pharmaceutical compositions, and use
IN Matthews, Ian Richard; Coulter, Thomas Stephen; Ghiron, Chiara; Brennan, Chris James; Uddin, Muhammed Kamal; Pettersson, Lars Olof Goeran; Da Graca Thirge, Dorthie; Huxley, Philip

PA Active Biotech AB, Swed.

SO PCT Int. Appl., 55 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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PI	WO 2004048378	A1	20040610	WO 2003-SE1805	20031121
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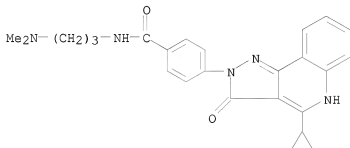
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OS MARPAT 141:38606				
IT 702704-93-6P,				
N-(3-Dimethylaminopropyl)-4-(4-cyclopropyl-3-oxo-3,5-				
dihydropyrazolo[4,3-c]quinolin-2-yl)benzamide 702705-30-4P,				
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N-(1-Benzyl-4-piperidinyl)-4-(4-cyclopropyl-6-fluoro-3-oxo-3,5-				
dihydropyrazolo[4,3-c]quinolin-2-yl)benzamide 702705-52-0P,				
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3,5-dihydropyrazolo[4,3-c]quinolin-2-yl)benzamide 702705-55-3P,				
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dihydropyrazolo[4,3-c]quinolin-2-yl)benzamide 702705-58-6P,				
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N-(2,2,6,6-Tetramethylpiperidin-4-yl)-4-(4-cyclopropyl-6-fluoro-3-oxo-3,5-				
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, N-[4-(Diethylamino)butyl]-4-(4-cyclopropyl-6-fluoro-3-oxo-3,5-				
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 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of pyrazoloquinolines and analogs as CD80 antagonists and immunomodulators)

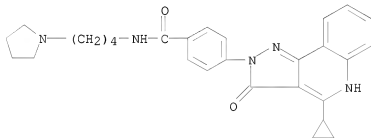
RN 702704-93-6 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[3-(dimethylamino)propyl]- (CA INDEX NAME)



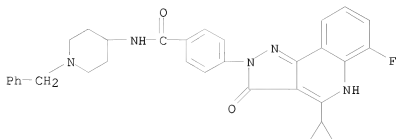
RN 702705-30-4 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[4-(1-pyrrolidinyl)butyl]- (CA INDEX NAME)



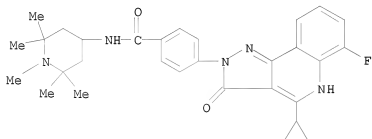
RN 702705-51-9 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[1-(phenylmethyl)-4-piperidinyl]- (CA INDEX NAME)



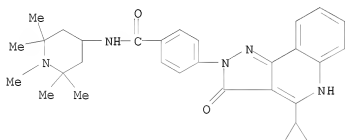
RN 702705-52-0 CAPLUS

CN Benamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(1,2,2,6,6-pentamethyl-4-piperidiny)- (CA INDEX NAME)



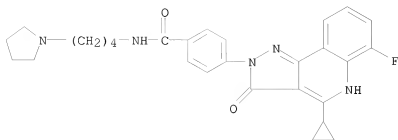
RN 702705-55-3 CAPLUS

CN Benamide, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(1,2,2,6,6-pentamethyl-4-piperidiny)- (CA INDEX NAME)



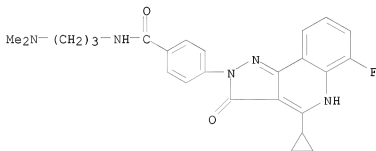
RN 702705-56-4 CAPLUS

CN Benamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(4-(1-pyrrolidinyl)butyl)- (CA INDEX NAME)



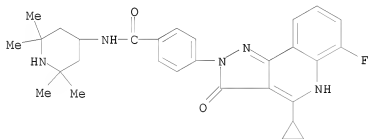
RN 702705-58-6 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[3-(dimethylamino)propyl]- (CA INDEX NAME)



RN 702705-59-7 CAPLUS

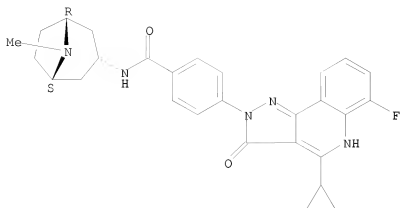
CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-(2,2,6,6-tetramethyl-4-piperidinyl)- (CA INDEX NAME)



RN 702705-60-0 CAPLUS

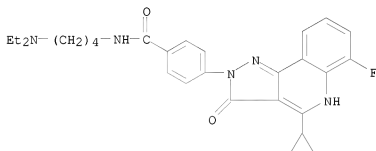
CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[(3-endo)-8-methyl-8-azabicyclo[3.2.1]oct-3-yl]- (CA INDEX NAME)

Relative stereochemistry.



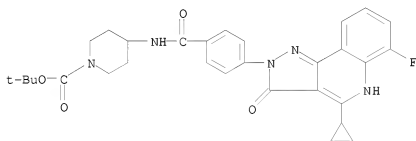
RN 702705-61-1 CAPLUS

CN Benamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[4-(diethylamino)butyl]- (CA INDEX NAME)



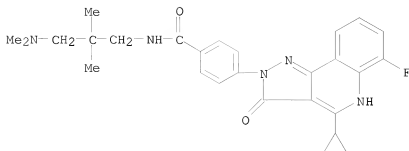
RN 702705-62-2 CAPLUS

CN 1-Piperidinecarboxylic acid, 4-[[4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)benzoyl]amino]-, 1,1-dimethylethyl ester (CA INDEX NAME)



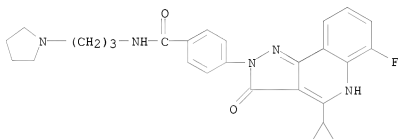
RN 702705-63-3 CAPLUS

CN Benamide, N-[3-[4-[(2-chloro-6-fluorophenyl)methyl]-1-piperazinyl]propyl]-4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-



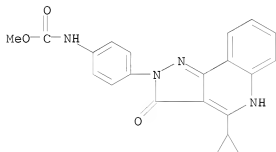
RN 702705-67-7 CAPLUS

CN Benzamide, 4-(4-cyclopropyl-6-fluoro-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)-N-[3-(1-pyrrolidinyl)propyl]- (CA INDEX NAME)



RN 702705-78-0 CAPLUS

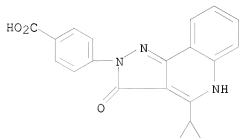
CN Carbamic acid, [4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)phenyl]-, methyl ester (9CI) (CA INDEX NAME)



IT 702706-05-6P, 4-(4-Cyclopropyl-3-oxo-3,5-dihydro-2H-pyrazolo[4,3-c]quinolin-2-yl)benzoic acid 702706-06-7P, 4-(4-Cyclopropyl-3-oxo-3,5-dihydro-2H-pyrazolo[4,3-c]quinolin-2-yl)benzoyl chloride
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (intermediate; preparation of pyrazoloquinolines and analogs as CD80 antagonists and immunomodulators)

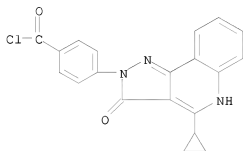
RN 702706-05-6 CAPLUS

CN Benzoic acid, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)- (CA INDEX NAME)



RN 702706-06-7 CAPLUS

CN Benzoyl chloride, 4-(4-cyclopropyl-3,5-dihydro-3-oxo-2H-pyrazolo[4,3-c]quinolin-2-yl)- (CA INDEX NAME)



RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT